

WHAT IS CLAIMED IS:

1. A purified and isolated polypeptide having part or all of the primary structural conformation and one or more of the biological properties of naturally-occurring pluripotent granulocyte colony-stimulating factor and characterized by being the product of procaryotic or eucaryotic expression of an exogenous DNA sequence.
2. A polypeptide according to claim 1 further characterized by being free of association with any mammalian protein.
3. A polypeptide according to claim 1 wherein the exogenous DNA sequence is a cDNA sequence.
4. A polypeptide according to claim 1 wherein the exogenous DNA sequence is a manufactured DNA sequence.
5. A polypeptide according to claim 1 wherein the exogenous DNA sequence is a genomic DNA sequence.
6. A polypeptide according to claim 1 wherein the exogenous DNA sequence is carried on an autonomously replicating DNA plasmid or viral vector.
7. A polypeptide according to claim 1 possessing part or all of the primary structural conformation of human pluripotent granulocyte colony-stimulating factor as set forth in Table VII or any naturally occurring allelic variant thereof.

8. A polypeptide according to claim 1 which has the immunological properties of naturally-occurring pluripotent granulocyte colony-stimulating factor.

5 9. A polypeptide according to claim 1 which has the in vitro biological activity of naturally-occurring pluripotent granulocyte colony-stimulating factor.

10 10. A polypeptide according to claim 1 further characterized by being covalently associated with a detectable label substance.

15 11. A polypeptide according to claim 10 wherein said detectable label is a radiolabel.

12. A DNA sequence for use in securing expression in a procaryotic or eucaryotic host cell of a polypeptide product having at least a part of the
20 primary structural conformation and one or more of the biological properties of naturally-occurring pluripotent granulocyte colony-stimulating factor, said DNA sequence selected from among:

25 (a) the DNA sequence set out in Tables VII and VII or their complimentary strands;

 (b) DNA sequences which hybridize to the DNA sequences defined in (a) or fragments thereof; and

 (c) DNA sequences which, but for the degeneracy of the genetic code, would hybridize to the
30 DNA sequences defined in (a) and (b).

13. A procaryotic or eucaryotic host cell transformed or transfected with a DNA sequence according to claim 12 in a manner allowing the host cell to
35 express said polypeptide product.

14. A polypeptide product of the expression of a DNA sequence of claim 12 in a procaryotic or eucaryotic host.

5 15. A purified and isolated DNA sequence coding for procaryotic or eucaryotic host expression of a polypeptide having part or all of the primary structural conformation and one or more of the biological properties of pluripotent granulocyte colony-stimulating
10 factor.

16. A cDNA sequence according to claim 15.

15 16. 17. A genomic DNA sequence according to claim

18. A manufactured DNA sequence according to claim 15.

20 19. A manufactured DNA sequence according to claim 18 and including one or more codons preferred for expression in E. coli cells.

25 20. A manufactured DNA sequence according to claim 18, and coding for expression of human species pluripotent granulocyte colony-stimulating factor.

30 21. A manufactured DNA sequence according to claim 20 and including one or more codons preferred for expression in yeast cells.

35 22. A human species pluripotent granulocyte colony-stimulating factor-coding DNA sequence according to claims 16, 17 or 18.

23. A DNA sequence according to claim 15 covalently associated with a detectable label substance.

24. A DNA sequence according to claim 23
5 wherein the detectable label is a radiolabel.

25. A single-stranded DNA sequence according to claim 23.

10 26. A DNA sequence coding for a polypeptide fragment or polypeptide analog of naturally-occurring pluripotent granulocyte colony-stimulating factor.

27. A DNA sequence coding for [Ala¹] hpG-CSF.
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28. A biologically functional plasmid or viral DNA vector including a DNA sequence according to one of claims 12, 15 or 26.

20 29. A procaryotic or eucaryotic host cell stably transformed or transfected with a DNA vector according to claim 28.

30. A polypeptide product of the expression
25 in a procaryotic or eucaryotic host cell of a DNA sequence according to claims 15 or 26.

31. A synthetic polypeptide having part or all of the amino acid sequence as set forth in Table VII
30 and having one or more of the in vitro biological activities of naturally-occurring pluripotent granulocyte colony-stimulating factor.

32. A synthetic polypeptide having part or
35 all of the secondary conformation of part or all of the amino acid sequence set forth in Table VII and having a

biological property of naturally-occurring human pluripotent granulocyte colony-stimulating factor.

33. A process for the production of a polypeptide having part or all of the primary structural conformation and one or more of the biological properties of naturally occurring pluripotent granulocyte colony-stimulating factor, said process comprising:
growing, under suitable nutrient conditions,
procaryotic or eucaryotic host cells transformed or transfected with a DNA vector according to claim 28, and isolating desired polypeptide products of the expression of DNA sequences in said vector.

34. Purified and isolated human pluripotent granulocyte colony-stimulating factor free of association with any human protein in glycosylated or non-glycosylated form.

35. A pharmaceutical composition comprising an effective amount of a polypeptide according to claims 1 or 34 and a pharmaceutically acceptable diluent, adjuvant or carrier.

36. A method for providing hematopoietic therapy to a mammal comprising administering an effective amount of a polypeptide according to claims 1 or 34.

37. A method for arresting proliferation of leukemic cells comprising administering an effective amount of a polypeptide according to claims 1 or 34.

38. A DNA sequence coding for
[Ser¹⁷]hpG-CSF.

39. A polypeptide product of the expression in a procaryotic or eucaryotic host cell of a DNA sequence according to claim 38.

5 40. A biologically functional plasmid or viral DNA vector including a DNA sequence according to claim 38.

10 41. A procaryotic or eucaryotic host cell stably transformed or transfected with a DNA vector according to claim 40.

15 42. A DNA sequence coding for an analog of hpG-CSF selected from the group consisting of:
[Ala¹]hpG-CSF;
[Ser³⁶]hpG-CSF;
[Ser⁴²]hpG-CSF;
[Ser⁶⁴]hpG-CSF;
[Ser⁷⁴]hpG-CSF;
20 [Met⁻¹, Ser¹⁷]hpG-CSF;
[Met⁻¹, Ser³⁶]hpG-CSF;
[Met⁻¹, Ser⁴²]hpG-CSF;
[Met⁻¹, Ser⁶⁴]hpG-CSF; and
[Met⁻¹, Ser⁷⁴]hpG-CSF.

25 43. A polypeptide product of the expression in a procaryotic or eucaryotic host cell of a DNA sequence according to claim 42.

30 44. A preparation of hpG-CSF which is greater than 95% pure and which comprises less than 0.5 ng of pyrogen per 0.5 mg of hpG-CSF.

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